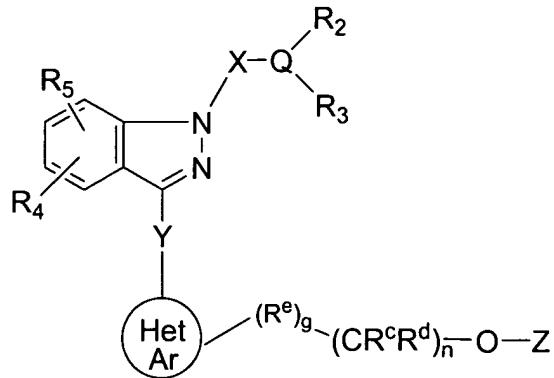


In the Claims

1 (Currently Amended) A compound of the structural formula I:



5

Formula I

or a pharmaceutically acceptable salt, *in vivo* hydrolysable ester, enantiomer, diastereomer or mixture thereof: wherein,

10 R represents hydrogen, or C1-6 alkyl;

R<sup>c</sup> and R<sup>d</sup> independently represents hydrogen or halo;

15 R<sup>e</sup> represents N or O;

X represents -(CHR7)<sub>p</sub>-, -(CHR7)<sub>p</sub>CO-;

Y represents -CO(CH<sub>2</sub>)<sub>n</sub>-, CH<sub>2</sub>, or -CH(OR)-;

20 Q represents N, or O, wherein R<sub>2</sub> is absent when Q is O;

R<sub>w</sub> represents H, C1-6 alkyl, -C(O)C1-6 alkyl, -C(O)OC1-6 alkyl, -SO<sub>2</sub>N(R)2, -SO<sub>2</sub>C1-6 alkyl, -SO<sub>2</sub>C<sub>6-10</sub> aryl, NO<sub>2</sub>, CN or -C(O)N(R)2;

25 R<sub>2</sub> represents hydrogen, C1-10 alkyl, OH, C<sub>2-6</sub> alkenyl, C1-6 alkylSR, -(CH<sub>2</sub>)<sub>n</sub>O(CH<sub>2</sub>)<sub>m</sub>OR, -(CH<sub>2</sub>)<sub>n</sub>C1-6 alkoxy, -(CH<sub>2</sub>)<sub>n</sub>C<sub>3-8</sub> cycloalkyl, -(CH<sub>2</sub>)<sub>n</sub>C<sub>3-10</sub> heterocyclyl, -N(R)<sub>2</sub>, -COOR, or -(CH<sub>2</sub>)<sub>n</sub>C<sub>6-10</sub> aryl, said alkyl, heterocyclyl, or aryl optionally substituted with 1-3 groups selected from Ra;

R<sub>3</sub> represents hydrogen, C<sub>1-10</sub> alkyl, -(CH<sub>2</sub>)<sub>n</sub>C<sub>3-8</sub> cycloalkyl, -(CH<sub>2</sub>)<sub>n</sub>C<sub>3-10</sub> heterocycll, -(CH<sub>2</sub>)<sub>n</sub>COOR, -(CH<sub>2</sub>)<sub>n</sub>C<sub>6-10</sub> aryl, -(CH<sub>2</sub>)<sub>n</sub>NHR<sub>8</sub>, -(CH<sub>2</sub>)<sub>n</sub>N(R)<sub>2</sub>, -(CH<sub>2</sub>)<sub>n</sub>N(R<sub>8</sub>)<sub>2</sub>, -(CH<sub>2</sub>)<sub>n</sub>NHCOOR, -(CH<sub>2</sub>)<sub>n</sub>N(R<sub>8</sub>)CO<sub>2</sub>R, -(CH<sub>2</sub>)<sub>n</sub>N(R<sub>8</sub>)COR, -

5 (CH<sub>2</sub>)<sub>n</sub>NHCOR, -(CH<sub>2</sub>)<sub>n</sub>CONH(R<sub>8</sub>), aryl, -(CH<sub>2</sub>)<sub>n</sub>C<sub>1-6</sub> alkoxy, CF<sub>3</sub>, -(CH<sub>2</sub>)<sub>n</sub>SO<sub>2</sub>R, -(CH<sub>2</sub>)<sub>n</sub>SO<sub>2</sub>N(R)<sub>2</sub>, -(CH<sub>2</sub>)<sub>n</sub>CON(R)<sub>2</sub>, -(CH<sub>2</sub>)<sub>n</sub>CONHC(R)<sub>3</sub>, -(CH<sub>2</sub>)<sub>n</sub>CONHC(R)<sub>2</sub>CO<sub>2</sub>R, -(CH<sub>2</sub>)<sub>n</sub>COR<sub>8</sub>, nitro, cyano or halogen, said alkyl, alkoxy, heterocycll, or aryl optionally substituted with 1-3 groups of R<sup>a</sup>;

10 or, R<sub>2</sub> and R<sub>3</sub> taken together with the intervening Q form a 3-10 membered carbocyclic or heterocyclic carbon ring optionally interrupted by 1-2 atoms of O, S, C(O) or NR, and optionally having 1-4 double bonds, and optionally substituted by 1-3 groups selected from R<sup>a</sup>;

15 R<sub>4</sub> and R<sub>5</sub> independently represent hydrogen, C<sub>1-6</sub> alkoxy, OH, C<sub>1-6</sub> alkyl, COOR, SO<sub>3</sub>H, -O(CH<sub>2</sub>)<sub>n</sub>N(R)<sub>2</sub>, -O(CH<sub>2</sub>)<sub>n</sub>CO<sub>2</sub>R, -OPO(OH)<sub>2</sub>, CF<sub>3</sub>, OCF<sub>3</sub>, -N(R)<sub>2</sub>, nitro, cyano, C<sub>1-6</sub> alkylamino, or halogen;

20  represents C<sub>6-10</sub> aryl or C<sub>3-10</sub> heterocyclyl phenyl, napthyl, phenanthrenyl, pyridyl, said aryl or heterocyclyl phenyl, napthyl, phenanthrenyl, pyridyl optionally substituted with 1-3 groups selected from R<sup>a</sup>;

Z represents (CH<sub>2</sub>)<sub>n</sub>PO(OR)(OR\*);

25 R\* represents hydrogen, or C<sub>1-6</sub> alkyl;

R<sub>7</sub> represents hydrogen, C<sub>1-6</sub> alkyl, -(CH<sub>2</sub>)<sub>n</sub>COOR or -(CH<sub>2</sub>)<sub>n</sub>N(R)<sub>2</sub>,

30 R<sub>8</sub> represents -(CH<sub>2</sub>)<sub>n</sub>C<sub>3-8</sub> cycloalkyl, -(CH<sub>2</sub>)<sub>n</sub> 3-10 heterocycll, C<sub>1-6</sub> alkoxy or -(CH<sub>2</sub>)<sub>n</sub>C<sub>5-10</sub> heteroaryl, -(CH<sub>2</sub>)<sub>n</sub>C<sub>6-10</sub> aryl said heterocycll, aryl or heteroaryl optionally substituted with 1-3 groups selected from R<sup>a</sup>;

R<sup>a</sup> represents F, Cl, Br, I, CF<sub>3</sub>, N(R)<sub>2</sub>, NO<sub>2</sub>, CN, -COR<sub>8</sub>, -CONHR<sub>8</sub>, -CON(R<sub>8</sub>)<sub>2</sub>, -O(CH<sub>2</sub>)<sub>n</sub>COOR, -NH(CH<sub>2</sub>)<sub>n</sub>OR, -COOR, -OCF<sub>3</sub>, -NHCOR, -SO<sub>2</sub>R, -SO<sub>2</sub>NR<sub>2</sub>, -SR, (C<sub>1-C<sub>6</sub></sub> alkyl)O-, -(CH<sub>2</sub>)<sub>n</sub>O(CH<sub>2</sub>)<sub>m</sub>OR, -(CH<sub>2</sub>)<sub>n</sub>C<sub>1-6</sub> alkoxy, (aryl)O-, -(CH<sub>2</sub>)<sub>n</sub>OH, (C<sub>1-C<sub>6</sub></sub> alkyl)S(O)<sub>m</sub>-, H<sub>2</sub>N-C(NH)-, (C<sub>1-C<sub>6</sub></sub> alkyl)C(O)-, (C<sub>1-C<sub>6</sub></sub> alkyl)OC(O)NH-, -

(C<sub>1</sub>-C<sub>6</sub> alkyl)NR<sub>w</sub>(CH<sub>2</sub>)<sub>n</sub>C<sub>3</sub>-10 heterocycll-R<sub>w</sub>, -(C<sub>1</sub>-C<sub>6</sub> alkyl)O(CH<sub>2</sub>)<sub>n</sub>C<sub>3</sub>-10 heterocycll-R<sub>w</sub>, -(C<sub>1</sub>-C<sub>6</sub> alkyl)S(CH<sub>2</sub>)<sub>n</sub>C<sub>3</sub>-10 heterocycll-R<sub>w</sub>, -(C<sub>1</sub>-C<sub>6</sub> alkyl)-C<sub>3</sub>-10 heterocycll-R<sub>w</sub>, -(CH<sub>2</sub>)<sub>n</sub>-Z<sup>1</sup>-C(=Z<sup>2</sup>)N(R)<sub>2</sub>, -(C<sub>2</sub>-6 alkenyl)NR<sub>w</sub>(CH<sub>2</sub>)<sub>n</sub>C<sub>3</sub>-10 heterocycll-R<sub>w</sub>, -(C<sub>2</sub>-6 alkenyl)O(CH<sub>2</sub>)<sub>n</sub>C<sub>3</sub>-10 heterocycll-R<sub>w</sub>, -(C<sub>2</sub>-6 alkenyl)S(CH<sub>2</sub>)<sub>n</sub>C<sub>3</sub>-10 heterocycll-R<sub>w</sub>, -(C<sub>2</sub>-6 alkenyl)-C<sub>3</sub>-10 heterocycll-R<sub>w</sub>, -(C<sub>2</sub>-6 alkenyl)-Z<sup>1</sup>-C(=Z<sup>2</sup>)N(R)<sub>2</sub>, -(CH<sub>2</sub>)<sub>n</sub>SO<sub>2</sub>R, -(CH<sub>2</sub>)<sub>n</sub>SO<sub>3</sub>H, -(CH<sub>2</sub>)<sub>n</sub>PO(OR)<sub>2</sub>, C<sub>3</sub>-10cycloalkyl, C<sub>6</sub>-10 aryl, C<sub>3</sub>-10 heterocycll, C<sub>2</sub>-6 alkenyl, and C<sub>1</sub>-C<sub>10</sub> alkyl,  
5 said alkyl, alkenyl, alkoxy, heterocycll and aryl optionally substituted with 1-3 groups selected from C<sub>1</sub>-C<sub>6</sub> alkyl, CN, NO<sub>2</sub>, OH, CON(R)<sub>2</sub> and COOR;  
10 Z<sup>1</sup> and Z<sup>2</sup> independently represents NR<sub>w</sub>, O, CH<sub>2</sub>, or S;  
g is 0-1;  
m is 0-3;  
n is 0-3; and  
15 p is 0-3.

20 2(Original). The compound according claim 1 wherein p is 1-3, Y is -CO(CH<sub>2</sub>)<sub>n</sub>, Q is N, X is -(CHR<sub>7</sub>)<sub>p</sub>, or -(CHR<sub>7</sub>)<sub>p</sub>CO-,.

25 3(Original). The compound according claim 1 wherein Q is O and R<sub>2</sub> is absent.

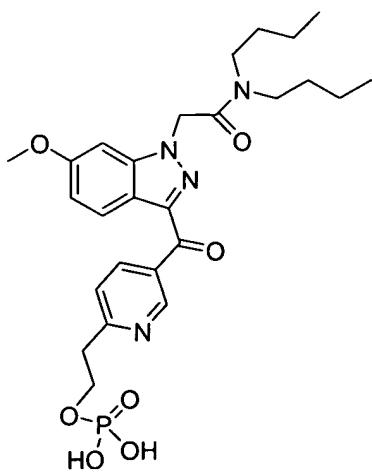
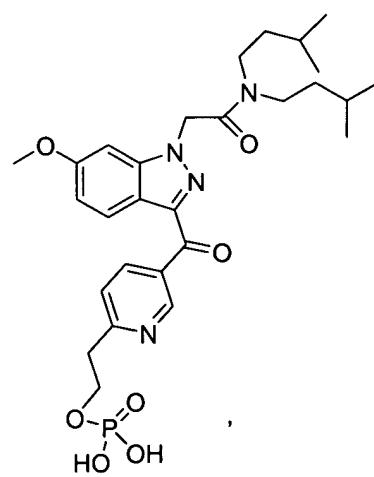
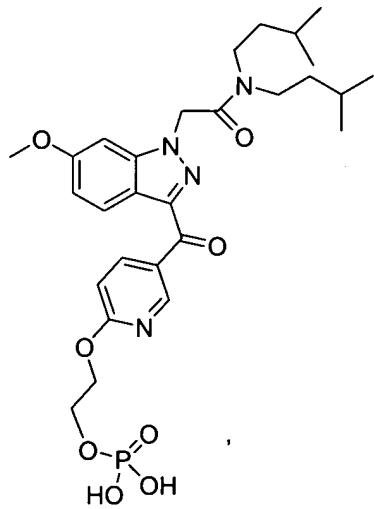
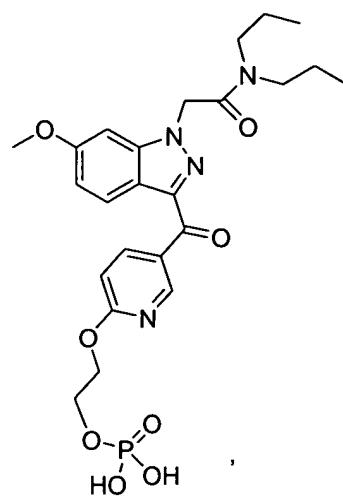
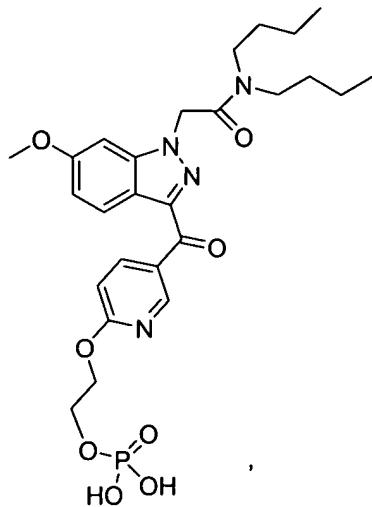
4(Original). The compound according to claim 2 wherein Z is PO(OR)(OR\*), R<sub>2</sub> is C<sub>1</sub>-10 alkyl or C<sub>1</sub>-6 alkylOH, Y is -CO(CH<sub>2</sub>)<sub>n</sub> and R<sub>3</sub> is (CH<sub>2</sub>)<sub>n</sub>C<sub>3</sub>-10 heterocycll, said heterocycll and alkyl optionally substituted with 1 to 25 3 groups of Ra.

30 5(Original). The compound according to claim 4 wherein  is a 6 membered heteroaryl or phenyl optionally substituted with 1-3 groups selected from Ra.

6(Currently Amended). A compound according to claim 1 5  
wherein  is pyridyl optionally substituted with 1-3 groups selected from Ra.

7(Original). A compound according to claim 1 which is in the form of a sodium or disodium salt.

8(Original). A compound which is:



or a pharmaceutically acceptable salt, in vivo hydrolysable ester, enantiomer, diastereomer or mixture thereof.

5                   9(Previously Presented).     A method for the treatment of ocular hypertension or glaucoma comprising administering a compound of formula I accordingly to claim 1.

10                  10(Previously Presented).   A method for the treatment of macular edema, macular degeneration, increasing retinal and optic nerve head blood velocity, increasing retinal and optic nerve oxygen tension, and/or a neuroprotective effect comprising administering a compound of formula I accordingly to claim 1.

15                  11.      Canceled.

12.      Canceled.

13(Original). A composition comprising a compound of formula I of claim 1 and a pharmaceutically acceptable carrier.

20                  14(Original). The composition according to Claim 13 wherein the compound of formula I is applied as a topical formulation, said topical formulation administered as a solution or suspension and optionally containing xanthan gum or gellan gum.

25                  15(Currently Amended). A composition according to claim 14 wherein one or more of an active ingredient belonging to the group consisting of:  $\beta$ -adrenergic blocking agent, parasympatho-mimetic agent, sympathomimetic agent, carbonic anhydrase inhibitor, EP4 agonist, a prostaglandin ~~or derivative thereof~~, hypotensive lipid, neuroprotectant, and/or 5-HT2 receptor agonist is optionally added.

30                  16(Original). A composition according to claim 15 wherein the  $\beta$ -adrenergic blocking agent is timolol, betaxolol, levobetaxolol, carteolol, or levobunolol; the parasympathomimetic agent is pilocarpine; the sympathomimetic agent is epinephrine, brimonidine, iopidine, clonidine, or para-aminoclonidine, the carbonic anhydrase inhibitor is dorzolamide, acetazolamide, metazolamide or brinzolamide; the prostaglandin is latanoprost, travaprost, unoprostone, rescula, or

S1033, the hypotensive lipid is lumigan, the neuroprotectant is eliprodil, R-eliprodil or memantine; and the 5-HT2 receptor agonist is 1-(2-aminopropyl)-3-methyl-1H-imdazol-6-ol fumarate or 2-(3-chloro-6-methoxy-indazol-1-yl)-1-methyl-ethylamine.